### REMARKS

#### Formal Matters

Claims 1-33 were pending in the application and were restricted into eight groups. Claims 1-11, 15, 19-30 and 32 are canceled. As discussed below, Applicants elect with traverse claims 12-14, 16-18, 31, and 33 (Group IV). Claims 12-14, 16, 31 and 33 are amended and claims 34-38 are added. No new matter is added by the amendments to the claims.

Support for the amendments is found throughout the specification such as at, for example, page 11, lines 23-25; page 11, line 27 to page 12, line 7; page 12, line 23 to page 13, line 2; page 13, lines 3-21; page 19, lines 22-23 (legend to Fig. 8) and Fig. 8; page 20, lines 17-22; page 20, line 15 to page 23, line 17; page 97, line 14 to page 98, line 7 and Fig. 4; and page 100, lines 5-10, Fig. 8, and Table 5 (page 100). No new matter is added by the amendments to the claims.

# Sequence Rules

Applicants are required to comply with the sequence rules as set forth in 37 C.F.R. § 1.821-25 at the time of election of the restriction. Applicants believe that they have already complied with 37 C.F.R. § 1.821-25 by submitting a Letter and Request to Use Computer-Feadable Sequence Listing Under 37 CFR § 1.821(e) upon filing the instant application on March 7, 2000 (copy of Letter enclosed). Specifically, the Letter requested that the computer-readable Sequence Listing filed in parent application Serial No. 09/070,416 be used as the computer-readable Sequence Listing for the instant application. A paper copy of the Sequence Listing and a statement that It is identical to the computer-readable copy from Serial No. 09/070,416 under 37 CFR § 1.821(e) was submitted with the Letter.

Applicants herewith submit another copy of the Sequence Listing for the systemic of the Evanises and in the unlikely sign metable that the

or gaver recommendation of the character along the Application of the commendation of the commendation of the character of th

Applicants submit a copy of the Letter and Request and 37 CFR § 1.821(e), another paper copy of the Sequence Listing, and state that the computer readable copy and paper copy are identical, that no new matter is added by the amendment. As a result, Applicants believe that they have complied with the Notice to Comply.

### Correction of Inadvertent Omission from the Specification When Filed

Upon filing the instant specification as a continuation of parent application 09/070,416 and provisional application serial no. 60/050,661, Appendix I was inadvertently omitted. Appendix I is a 15-page table comparing sequence identities between various light chain sequences. Insertion of the table into the specification does not add new matter because the table was present in the provisional application 60/050,661 to which the present application ultimately claims priority. Insertion of the Appendix as Table 6.1-6.15 is respectfully requested.

In a related application, U.S. application serial no. 08/850,058, the position of Appendix I was objected to. Appendix I was after "What is claimed is:" on page 103 and before the Claims. Applicants renamed the table as Table 6.1-6.15 and repositioned it to immediately before "What is claimed is:". Applicants respectfully offer this positioning scheme for consideration in the instant application.

Applicants submit Table 6.1-6.15 on fifteen pages. The word "Appendix" and the original page number on each page of the original appendix are deleted and "Table 6.X" is inserted therefor, where "X" refers to subpart 1-15 of Table 6.

The word "Appendix" occurs only once in the originally filed specification at page 96, line 24. The word "Appendix" has been deleted from the specification and the term "Table 6.1-6.15" has been inserted therefor.

: 8

" now haven has been alled by these amendments to the

# Other Amendments to the Specification

Applicants amended the specification on page 13, line 27 to correct a typographical error by replacing a semicolon (";") with a period ("."). No new matter was added by the amendment to the specification.

Applicants amended the specification to correct a typographical error in the legend for Fig. 2A-2C on page 17, line 30. Specifically, Fig. 2C shows the sequence of a portion of the nucleic acid construct depicted in Fig. 2B. Originally filed Fig. 2C indicates that that sequence is SEQ ID NO:13 and thus provides support for the amendment. Correction of the legend for Fig. 2C has been corrected accordingly. No new matter has been added by the amendment to the specification.

## Election/Restriction

The Examiner has indicated that the application contains claims directed to patentably distinct species of the claimed invention and requires restriction under 35 U.S.C. § 121 according to the eight groupings indicated in the Office Action (Paper No. 4, mailed July 5, 2001)

In addition, Applicants were asked to further elect patentably distinct species of the claimed invention as indicated on page 4 of the Office Action. Applicants elect with traverse, for the reasons stated herein, the following species:

The constant domain is from a human IgG.

The anti-Ob-R'anti-HER3 species.

Applicant respectfully traverses the restriction and election requirement as applied to the currently pending claims for the reasons provided below.

Applicant respectfully traverses the restriction requirement in which free thicl for protuberance/cavity structures within the multimentant of implication of the support the restriction. The

under de la company de la Entrenta de la company de class 530 and subclass 387.3 as another multispecific antibody of the invention comprising a protuberance and cavity in the multimerization domain (Group IV). Thus, the Examiner would not be placed under an undue burden to search in the same class and subclass of art in order to evaluate the patentability of claims in Groups III and IV, for example. The same argument applies to Groups I and II, Groups V and VI, and Groups VII and VIII. Applicants respectfully submit that the restrictions that distinguish Group I from Group II, Group III from Group IV, Group V from Group IV, and Group IV from Group VIII should be withdrawn.

Without acquiescing to the restrictions, however, and merely to expedite prosecution of the claims, Applicants elect with traverse Group IV, Claims 12-14, 16-18, 31 and 33 drawn to a multispecific antibody, wherein the multimerization domain is altered to comprise a protuberance and a cavity.

With respect to the election of species requirement, Applicants respectfully traverse the election requirement for failing to recognize Applicant's right to allowed claims that link a reasonable number of species under 37 CFR § 1.141. It is Applicant's understanding and right under 37 CFR § 1.141 that, following election, the claims will be examined fully with respect to the elected species and further to the extent necessary to determine patentability for a reasonable number of species encompassed by the generic claims.

With the above reservation of right, Applicant elects, with traverse, a constant domain from a human IgG and further elects anti-Ob-R/anti-HER3, an illustrative example of the claimed multispecific antibody of the invention.

A marked-up version and a clean version of the pending claims is attached.

If the Examiner has any questions, the Examiner should feel free to

This document is timely filed with a petition and fees for a three-month extension of time. In the unlikely event that additional fees are due, Applicants hereby petition the Commissioner to authorize any extensions of time and/or to deduct fees from or add credits to our Deposit Account 07-0630 as necessary to maintain the pendency of this application.

Respectfully submitted,

GENENTECH, INC.

Date: November 5, 2001

3y: <u>/</u> ∠

Deirdre L. Conley, Ph.D.

Reg. No. 36,487

Telephone No. (650) 225-2066

09157

PATENT TRADEMARK OFFICE

Doc. #99911

### VERSION WITH MARKINGS TO SHOW CHANGES MADE

### In the Specification:

Deleted information is shown in strikethrough  $(\Theta)$ , and added information is shown as <u>underlined</u>.

Paragraph beginning at page 13, line 26, has been amended as follows:

(ii) recovering the multispecific antibody from the host cell culture;  $\underline{\cdot}$ 

Paragraph beginning at page 17, line 16, has been amended as follows:

Figs. 2A-2C. Fig. 2A diagrams a selection scheme for Ca3 heterodimer using phage display vector, pEA2. Phage displaying stable CG heterodimers are captured using an antibody directed to the gD flag. Fig. 2B diagrams a didistronic openon in which Ca3 expressed from a synthetic gene is co-secreted with a second copy of  $C_{\rm B}3$  expressed from the natural gene (Ellison et al. Nucleic Acids Res. 10:4071-4079 (1982)) as a fusion protein with M13 gene III protein. The synthetic  $C_93$  gene is preceded by a sequence encoding a peptide derived from herpes simplex virus glycoprotein D (dD flag, Lasky, L. A. and Dowbenko, D. J. (1984) DNA  $\underline{3}$ :23-23; Berman, P. W. et al., (1985) Science 327:1490-1492 and a cleavage (G) site for the site-specific protease, Generase : (Carter, F. et al. (1989) Proteins: Structure, Function and Genetics 6:240-248). Fig. 2C is the nucleic acid sequence of the dicistronic operon (SEQ ID-NO:1) (SEQ ID NC: 13) of Fig. 2B in which the residues in the translated 0.3 genes are numbered according to the

ingen i transporte de la companya de la companya de mendementant de la companya d

gene (366, 368, and 407).

Paragraph beginning at page 95, line 8, has been amended as follows:

--A large human single chain Fv (scFv) antibody library (Vaughan et al. (1996), supra) was panned for antibodies specific for eleven antigens including Axl(human receptor tyrosine kinase ECD), GCSF-E (human granulocyte colony stimulating factor receptor ECD), IgE (murine IgE), IgE-R (human IgE receptor  $\alpha$ chain), MPL (human thrombopoietin receptor tyrosine kinase ECD), Musk (human muscle specific receptor tyrosine kinase ECD), NpoR (human orphan receptor NpoR ECD), Ese (human receptor tyrosine kinase, Rse, ECD), HER3 (human receptor tyrosine kinase HEF3/cerbB3 ECD), Ob-R (human leptin receptor ECD), and VEGF (human vascular endothelial growth factor) where ECD refers to the extracellular domain. The nucleotide sequence data for scFv fragments from populations of antibodies raised to each antigen was translated to derive corresponding protein sequences. The V sequences were then compared using the program "align" with the algorithm of Feng and Doolittle (1985, 1987, 1990) to calculate the percentage identity between all pairwise combinations of chains (Feng, D.F. and Doolittle, R.F. (1985) J. Mol. Evol. <u>21</u>:112-123; Feng, D.F. and Doclittle, R.F. (1987) J. Mel. Evel. 25:351-360; and Feng, D.F. and Doolittle, R.F. (1990) Mothods Engymol. 183:375-387). The percent sequence identity results of each pairwise light chain amino acid sequence comparison were arranged in matrix format (Appendix) Table 6.1-6.15).

The Approvedix is amongled to be tope Table 6.1-6.15 as follows:

APPENDIX -1- Table

53 MusK.01 54 MusK.02 55 MusK.06 56 NpoR.25 57 NpoR.44 58 NpoR.53 59 NpoR.81 60 NpoR.86 50 Mpl.32 51 Mpl.33 52 Mpl.35 43 Mpl.21 44 Mpl.24 45 Mpl.26 46 Mpl.28 69 Rse.18
70 Rse.20
71 Rse.21
72 Rse.22
73 Rse.23
74 Rse.24
75 Rse.52
76 Rse.53
77 Rse.53
77 Rse.58
78 Rse.60
79 Rse.61 65 Rse.07 66 Rse.08 67 Rse.15 68 Rse.16 61 Rse.01 62 Rse.02 63 Rse.03 64 Rse.04 42 Mpl.19 41 Mpl.16 49 Mpl.31 48 Mpl.30 47 Mpl.29 82 her3.10 81 her3.1 80 Rse.63

APPENDER -2- 79607

111 vegf.10 112 vegf.2 113 vegf.3 114 vegf.4 115 vegf.5 101 obr.2 102 obr.20 103 obr.21 104 obr.22 105 obr.23 106 obr.24 107 obr.26 108 obr.3 109 obr.4 97 obr.16 98 obr.17 99 obr.18 100 obr.19 86 her3.18 87 her3.19 88 her3.22 83 her3.11 84 her3.12 85 her3.16 90 her3.4 91 her3.7 95 obr.14 96 obr.15 116 vegf.6 117 vegf.8 94 obr. 12 93 obr.11 92 obr.1 89 her3.3 110 vegf.1 ω 4 ഗ Ax X 6 æ 9 10 12 <u>1</u> 14 15 16 œ 19 20 22 23

APPENDIX 3 6.3

APPENDIX 4- Chy

_		_						<u> </u>	r		_						_			_		_		$\neg$		<sub>1</sub>		_	_	_		_		_	_	_			т		т	7
																	$\cdot \mid$	61	43	61	60	61	9		98		58	43	62	45	61	63		48	43	48	44	61	59	44	24	Ē.
																$\cdot \Big $	43		82	48	48	47	43	42	42	47	44	99	47	85	47	49	48	79	98	78	84	48		80	25	
	T											Ì			$\cdot$		100	61	43	61		61	001	91	98	62	58	43	62	45	61	63	58	48	43	48	44	61		44	26	
					1			Ì							61		61	75	45		67			57		76	76	48	74	48	80	64	62	52	48	52	46	88	66	48	27	9
					7										99	43		60	43		59		99			61	57	43	61	45	60	62	57	48	43	48	44		58	4	28	
	1		1		1		1						60	8	6.	47	61	94	45	73	72	9	61	58	59	90	95	47	85	48		66		52	47	52	46	73	71	49	29	
-	1	1	1	_	1				1		·	60	98	60	99	45	99	60	45	60	6	_	99	90	97	61	57	45	61	47	60	62	57	50	45	48	46		58	46	30	7
	†	7	1	_	7					•	87	57	87	55		43	88				59	57	88	97	87	58	54	43	56	45	57	58	55	47	43	45	42		57	44	31	
-	1									88	99	59	99	59	100	44	100	59	44	59	59	59	100	91	98	60	56	44	60	47	59	61	56	49	44	48		59	57	46	32	٦
-	+		$\dashv$						44	43	45	47	43	╗	4	<u></u>	7	49	8 1	48	48	4	4	42	42	47	44	99	47		47	48	48	78		78	83	48	47	79	33	
-	1	1						47	-1	67	75	79	7	8.5	7	47	72	74	47	68	68	79	72	66	71	76	75	47	74	49	79	64	62	53	47	53	48		66	50	34	
, }	$\dashv$	+		-		-	50	08	46	4	46	-	45		$\vdash$	$\dashv$		-	$\vdash$	-	Н	-	_	-	$\vdash$	$\vdash$	47	$\dashv$	Н	$\dashv$	50	50	47	83	79	83	81		50	100	35	1
DENINT	1					51	7	48		6	67	-	6	$\dashv$		$\vdash$	67		$\vdash$	$\vdash$	-	-	-		$\vdash$	$\vdash$	63	$\blacksquare$	Н		$\neg$			$\vdash$			50	77	76	5 1	36	
{	+	+				5	-+	47	$\vdash$	5	61	-	5	9	$\dashv$	$\vdash$	$\dashv$	_	Н		$\vdash$			-	_	-	$\vdash$	$\vdash$	-		-	_	$\vdash$	52	47	52	46		68	49	37	
`				47	4		47	94	44	43	4	4	4	4	44		_	_	8 1	-	$\vdash$	-			$\vdash$	-	45	-				1			9	7	8	49	48	79	38	
1			9	4	46	7	4	9		3	4	4		4	40	9	4	4	78	4	4	4	4	ယ	38	4	4	9	4	8		44		7	95	7	80	44	43	78	39	MP
1			8	4	5	8	4 8	83	45	4	4	4	4	4	4	8	-	4	9.8	4	4	$\vdash$	┝	┝	-	$\vdash$	44	$\vdash$	-	85	46	4	4	8	-	8	10	4	-	8 1	Н	<u>'</u>
	_	0 8	3 8:	6 4	0 5	1 8	$\square$			<b> </b>	3 44	_	$\square$	_	$\vdash$		_	-	<b>-</b>	<u> </u>	-	├─		⊢	-		┤─	$\vdash$	┢	_		-	-	├	<del> </del>		010	4	<del> </del> —	8	4	
	0	4	4	9	6	5	8	4	6	5	Н	8	6	9	6	4	9	7	_	7	6	<u></u>	6	S	6	7	79	4	7	4	8	6	6	5	4	5	4	70	6	4	4	H
-	8	9	7 9	4	4	8	4	9		4		4	1 4	4	2 4	9	4	4	æ	4	4	4	4	4	4	4		=	┪	8	4	4	4	7	9	7	84	4	4	-	4	
ŀ	_	6 9			_	_	_	_	_	_	5 4	_	_	_	_	<b>"</b>	4	(2)	æ	4	4	4	4	4	4	4	-	=	1	8	4	4	4	7	9	7	8	4	4	80	4	П
	4	4	Ι.					Ι.			14 6					_			_	_	_	<u></u>		(n	(n	_	۱,,	_	_	_	6	6		(n	_	(D	4		7	(n	7	1 (
1	71	4	u. Ur			t : Hillo	,	ر ديا ا	† : +-	+		1		Ιω	•	¶ 1 - 2-			5 , 7	T . ++	7 . ++	•-		T		1	1	I L	4.	! . :		!			-	-		1		0		!!
i	-	-2-	4	6	7	4	6	4	5	5	S S	_ _		1 .	Çn	4	cn	6	4	İo	6	0	ľ	5	G	i lo	l G	4	6	4	10								6.5	•	4	
	5	4	-	-	-	3	-	╁.		5	-	-	3 5	-	1,5		(7)						'n	(n	(D	1	7 6	_	1	_	6	7	6	4	4	4	4	7	17	4	7 48	
	4.	2 4	4.2	<u>б</u>	7	47	6	4	5	5	5	σ	5	6	6	12	6	_	4	7	7	6	6	5	5	6	6	4	5	4	σ,	7	8	4	47	12	4	1	7	5 4 (	4	H
	7 4	-	<del> </del> _	₩.		ļ.,	ļ.	-	-	╀	9 6	<del> </del>	↓_	<b>↓</b>	0	_ _	6	0	4	1	1	6	0	(n	(n	1-	6	4	0	4	6	7	7	4	4	5	7	-	1 0	4	5	
	7   4	ω	7	4	5	8	6	2	0	5	0	0	0	100		5	100	α ,	6	0	0	5		7 5	9	5		5	1	9	5 7	2 6	6	9	4	. U	17	6.	6	4	0 5	$\dashv$
l	4	6		ω	4	7	4	6	β	2	7	٢	0	9		c	7	٦	ω	Ū	الم	9		N	U	4	4	6	^	1	9	_	0	0	6	1-	4	U	ıω	17	1-	

)

APPENDIX 6-6

				_	1		_			$\neg$			_	_	$\neg$	一		$\neg$	Т	Т	Т	Т	1	Т	Т	Т	Т	Т	Т	Т	Т	Т	Т	T	٦
E E	24														_		$\downarrow$	_	1	1	1	$\downarrow$	1	1	1	1	$\downarrow$	1	$\downarrow$	$\downarrow$	$\downarrow$	$\downarrow$	4		4
	25																															_	╛		
	26																1																		
99		T		Ī													<u>.</u>																		
Ĩ	28															_	W.50.		1																
H	29	1	H				$\exists$									_	500.8	1	1	1	1	1	1	1	1	1	1			1		1	1		
Н	30	+		_		4										-	<del>(%</del>	+	+	$\dagger$	+	+	$\dagger$	+	+	+	+	+	1	+	1	-	1	_	7
H	0 31	+	H	$\dashv$				$\exists$	H	H			Н					1	1	$\dagger$	+	$\dagger$	$\dagger$	+	$\dagger$	1	$\dagger$	+	$\dagger$	1	1	$\dashv$	+	1	┨
H	⊢	+		-					_	-				_				$\dashv$	-	$\dashv$	$\dashv$	+	+	+	+	$\dagger$	+	1	$\dashv$	-	$\dashv$		$\dashv$	$\dashv$	$\dashv$
-	32 3	+							_	_	-	-		$\vdash$	_			$\dashv$	$\dashv$	$\dashv$	+	+	+	+	+	+	+	$\dashv$	-	$\dashv$	-		$\dashv$	-	$\dashv$
	33	-	-						-	-	-	_		-					$\dashv$	+	$\dashv$	+	+	$\dashv$	+	+	$\dashv$	-	$\dashv$	$\dashv$			$\dashv$	-	$\dashv$
	34	_	_		_		_		_	-		_		_		_		-		$\dashv$	$\dashv$	$\dashv$	4	$\dashv$	$\downarrow$	$\dashv$	$\dashv$	$\dashv$	$\dashv$	$\dashv$	$\dashv$			$\dashv$	$\dashv$
	35	_					_		L	_	_	_	_	_	_	_			4	_	$\dashv$	4	4	$\downarrow$	$\downarrow$	4	_	$\dashv$		$\dashv$					$\dashv$
	36													_	_	_						_	$\downarrow$	_	$\downarrow$				$\dashv$						4
	37												L				_																		
	38																															_			
MP	39																																		
	4																																		
	4	1							T		<u> </u>																								
$\vdash$	42	+		╁╴			$\vdash$	$\dagger$	T	1							-																		П
$\vdash$	43	+	+	$\vdash$	-	-		+	╁	$\dagger$	+	$\dagger$	+	+	$\dagger$	-	-						1							-					П
-	+-+	+	+	-	╁	$\vdash$	╀	+	╁	+	+	$\dagger$	+	+	+	+	+			-								_	-		-	$\vdash$	T	$\vdash$	H
-	44 4	-}-	+	$\vdash$	-	$\vdash$	-	+	╀	+	+	+	+	╁	+	+	-	$\vdash$	-	-								-	-	-	-	-	$\vdash$	-	H
	45 46	2.5			•	•	•	,	İ	•	,	•	'	1	'	ī	ı	1	!	1	ļ	,		' '	' '			!	1	1	f	1	ľ	Ī	! !
<u> </u>	157	-		+	+	+-	1	+	+-	+	_	<u> </u>	1	1-	1	1	1	1	1	<del></del>	1-								ţ	1		· 1	- 1		
	47	-	-		-	_	-	_	1	_	-	1	$\downarrow$	1	_	1	-	1	_	-	_	_	_		_			_	-	-	-	-	-	-	
	48	-		_	-			1		1	_	_		_	_	_	_	-	_	_	_	_	_		_		_	_	_	-	-	-	-	-	$\sqcup$
	49																			_	_		_		_	_		_	_	_	-		-	-	
	50																																		
- 1	ĮΥ		1	l	- 1	- 1	1	- 1		ŧ	- 1	- 1	- 1		- 1		. [	_}	1	1.	1_	1_	-	1_	↓_	-	-	1_	┷-		+-		-+-	+	+-

APPENDIX 7- /aski 7

[			94	47	4 8	79		98	43	42	44	47	42	48	4	98	42	49	& -1	48	48	47	42	4		47	44	99	47		47	48	48		98	77		48	47	79	52	ĺ
	8 8 4	9	95	4	4	<u></u>	4	99	4	43	4	4	-	4	┧	۵	-	7	┥			47	43	42	42	47	44	100	47				48	79	99				47		Musk 53	
	8	9	94	4	4	7	4	98	4	$\dashv$	4	4	-	4	4	┪	4	49	_	48	7	-	42	$\dashv$	4	4	_	۵	47	85	47	48	48	78	98	77	83		47	79	54	
	5	<u>_</u>	83	4	-	<u>_</u>	_	<u>_</u>	4	42	4	46	4	4	-1	8	┥	48	$\dashv$	$\dashv$	$\dashv$	46	44	42	42	48	4	84	45	85	46	49	48	83	83	82	9	49		8-1	55	
	<u>0</u> 81	7	75	5		<u>@</u>	5	76	47	4	$\dashv$	49		50		77	4	5	-	51	5	49	46	43	4		46	77	47	81	49			80	76	79	8	51	50	8 1	56	
	44	40	44			45		43	100	88		61	99	61	100	43	100	61		61	60	61	100	91	98		58		62	45	61	63	58		43	48	44	61	59	44	57	
Ì	48	43	47	46		46	48	46	4	48	48	48	49	4	4	╛	4	49	47	4.7	46	48	49	48			46	46	45	46	48	46	44	48	46	48	48		46	45	NpoH HodN	
	46	43	45	43	45	46	45	44	47	49	47	47		45		44	48	49	45	46	45	47	48	49			45	44	46	46					44		46	46	45	45	59	
	84	96	95		48		47	99	44	43	45	47	43	48		99	43	49		48	48	47	43	42	42		44	300	47	85			48		99		84	48		80	60	
	47	43	47		73		63	47	58	57	59	66	58	60	59	47	59	70	46	75		66		57		65	61	47	62		99	75	80	49	47	49	47	75	73	46	61	
	47	43	47	64		47	63	47	58		59	66	59	60		47	0.9	70	46	75	74	66	60	58	58	65		47	62	48	66		80	49	47		47	75	73	46	62	
API	46	43	48		65		73	47		56	59		59	74	0.9		0.9	89	45	73	72	89	60			88		47	80			66	65	51	47		46	73	71	46	63	
APPENDI	45	42	47		65			46		56	59			08	0.9	46	60	93		72	71	99	60	57		89		46	84	47		65		51	46	51	45	72	70	48	64	
1/	88	72	74	41	47	73	42	74	41	36	40	41		40	40		40		86		43	41	40	38	38	42	40	74	40	77	41	42	41	74	73	80	88	43	42	72	65	
7- /a	100	80	83	46	50	81	48	83	45	42	46	46	44	46	44	84		48	98	49		46	44	42	42	48		84	45	85	46	49	48		83		100	49	48	81	66	
2 fe	46	43	48			50	79	47		57			-	80	61	47	61		45		72	100	61	58		90		47		48	100	66		52	47		46	73	71	49	67	
4)	43	43	43	83	64	46	78	43		52			53	79	54	43	54		42		71	95		52	52	86	90	43	84	45	95	61	60	48	43		43	71		46	68	
	47	44	49	76	66	48	74	48	59	57	60	90	60	75	61	48	61	90	46	73	72	90	61	58	59	89	86	48	81	48		99		52	48			73	71	47	69	
	44	43	46		64	48	77	45	57	55	58	98	58	79	59	45	59	92	43	71	70	98	59	57	57	88	93	45	83	46	98	64	62	50	45	50	44	71	70	47	70 70	4
	45	44	47		65	46	73	46	58	56	59	88	58	73	59	46	59	88	44	72	72	88	59	57	57	88	86	46	<b>-</b>	6	8	5	4	<b>├</b> ─	6	0	<b>↓</b> —	2	70	45	71	1
	100		83		50	81	48		45	42		46	44	46	44	84	44	48	98	49	49	6	4	42	42	48	44	84	45	85	_	49	48	83	83	82	100	49	48	8 1	72	-
		43		84						57			60	80	61	47	6-	94	45	┼-	72	lo o	1-	+-	59		$\vdash$	⊢	╁╌	48	100	6	4	52		2	╀	ω	╁╾	49	73	-
			82							14-	101	4.5	ıω	15	<u>بدا</u> س	ω	(w		•	( <u>ئ</u> ى ,	i Œ	U1	•	IN	•	•	•	<u>ω</u>	4-	85		(d)	4.		100	œ r	12	14	+	ļ	-	
			50							60			63		64	9	4	3	Ü	15	4	ω	4	12	┼─	9	8	9	~	51	ω	4	8	5	9	5	9	15	83	<del> </del>	5	i
	49	44	49		74			48		57		<u>ω</u>	┼-	8	9	48	59		8	6	5	ω	9	100	╁	6	<del> </del>	8	0	50	ω	5	6	1-	8	12	+	6	4	9	76	1
	44	1		59	7	-	2	43	C	3 0	99	-	+-		0	۳		1.		6.1		6.1		+	8	62	8	ω	2	45		ω	8	48	ω	8	4	-	59		77	
	4/		47	64		47				5 7		00	59	60	60	4	60		46		14	66	60	5 8	58	65	61	47	2	48	6	75		49					73	46	78	-
	α-1		19		5	100	50		4		6	2	4		44		44		9	0	0	4 9		4	42	48	46	80	ů	90	49	49	46	83	19		8	50	49			

APPENDIX -9 6.9

	52	$\exists$	T	T		Ī				ľ		T							Γ																	$ \top $	
MusK		+	$\dagger$	+	$\dagger$	1	_			t	$\dagger$	+	_		_					-	$\vdash$			-												1	┪
٤	3 54	$\overline{}$	+	+	+	1	-		-	┞	$\dagger$	+	1		_	H	-	$\vdash$	-	$\vdash$		$\vdash$	$\vdash$													7	┪
$\vdash$	4 55	$\rightarrow$	$\dashv$	+	+	$\dashv$	$\dashv$		$\vdash$	+	+	+			_	-	$\vdash$	+		$\vdash$	$\vdash$	╁	H	$\vdash$												_	$\dashv$
-	ш	-	+	+	+	-	_		-	╀	+	+				-	H	+	-	L	-	-		├													-
-	56		+	+	$\dashv$	$\dashv$	-	_	-	┞	+	+	$\dashv$			$\vdash$	-	$\vdash$	+	-	-	-	$\vdash$	-				_									$\dashv$
Z	57	-	1	4	4	$\dashv$		_	-	L	+	4	4		_	-	$\vdash$	╀	-	$\vdash$	-	-	-		-	$\vdash$			_			_					-
Noon	58	_	4	$\downarrow$	$\dashv$	-		-	$\vdash$	╀	1	4	_		L	_	-	$\perp$	$\vdash$	H	-	-	-	-	-	_		$\vdash$	_			-		_			$\dashv$
_	59	$\vdash$	-	4	$\dashv$			_			1	4	_		-	-		$\downarrow$	-	-	-	-	L	-	-		_	Н	_			-		_			$\dashv$
L	00	-		_	_				L	ļ	1	_			_		L	1	1	L	_	_		-	_	_						_		_			
_	61		$\perp$	_	$\rfloor$			_		$\downarrow$	1	_			_			1	*			1	L		_	_	_		L	_		_		_	_		Н
	62									l		$\downarrow$		L		L	_	_	-	L		L	L	_						_	_	_	_		_		
	63																		\\ \ \ \ \ \ \ \ \ \ \ \ \ \ \ \ \ \ \	L				_		_	_		_	_	_		_	_	_		
	64																		Ŀ													_		_			Ц
	65																													_				_	_		
	66																																				
	67										1	1							Ţ.																		
	68			1						Ť																											
	69	1-1		1					1	1							T				T	T															
#88 H	₩	┥		1			-			t	1			$\vdash$	-	1	$\dagger$	$\dagger$	  -  -			T	<b>†</b>	1		-											
	2	-		1		_	-	1	+	$\dagger$	1		_	-		T	$\dagger$	†			1	1	1	T		T	<del> </del>		T			1					П
-	72	1						t	$\dagger$	$\dagger$	1				$\vdash$	1	t	$\dagger$	3.		$\dagger$	+	T	$\dagger$					T	T		<u> </u>	T	T			
-	+-	-		_		-	-	+	+	$\dagger$	-	_	-	$\vdash$	-	†	$\dagger$	$\dagger$			$\dagger$	$\dagger$	†	+	$\dagger$	$\dagger$	-	$\vdash$		1	-	T		$\dagger$			H
,	73 -4	1	!			,	7		ļ	Į	į			!	!	!	•	!	1		!	!	!	!		!	!	!	ļ	!	!	Ĩ	!	!	!	!	!
ı.	10	1				· · · ·	1	1	1	1				Ī	 !		Ì		1		i		1		<u>.</u>	1	!		i	i	ļ		1	İ			
-	1	1				-	$\vdash$	+	+	$\frac{1}{1}$	-		-	╀	+	+	+	+	┽-		+	+	+	+	+	+	-	-	╁	+	-	+	╁	+	+	+	$\vdash$
-	ō	<del>†-</del>			-	_	-	+	+	+	_		-	-	+	+	+	+	+	+	+	+	+	+	+	+	-	+	-	+	+	+	+	+	+	-	-
-		<del>1</del> –	_		_		-	-	-				-	-	-	-	-	-	$\downarrow$	+	-	+	-	$\downarrow$	+	+	-	-	-	-	-	+	-	-	-	1	
	à	1-	_	_	_	-	-	+	1		_		-	1	-	+	1		1	-	-	-	-	1	-	+	-	-	1	+	+	$\downarrow$	+	_	-	+	-
	۵	1																		1																	

APPENDIX =10- 6/248

	46	42	46	63	72	46	62	46	57	56	58	65	58	60	59	46	59	70	45	74	73	65	59	57	57	64	60	46	61	47	65	74	80	48	46	48	46	74	72	45	80	
	00		83	46	50	81	48	83	45	42	46	46	44	46	44	84	44	48	98	49	49	46	44	42	42	48		84	45	85	46	49		83		82	100		48	81	81	
- 1	<u>ω</u>	78	79	49	51	100	50	79	46	44	46	49	44	48	4	80	4	49	79	50		49	44	43	42	48	46	0.8	45		49			83		83	_	50	49	100	82	
			84	49	51	85	49	85	46	46	47	49	45	49	45	86	45	51	83	52	52	49	45	44	43	49	46	86	47		49				85		85	52	51	85	83	
	49		50	71		51	72		63	61	64	76	64	69	65	49	65	76	48	94	93	76	65	62	63	75	71	49	70	51		80			49	54	49		92	51	84	
		43			64		84	47	58	54	59	0.8	59	99	09	47	60	74	44	67	99	0.8	60	56	88	75	75	47	73	47	80	63	61	51	47		45	7	65		85	
		96	95	47	48	08	47	99	44	43	45	47	43	48	43	99	<u>Δ</u>	49	82	48	48	47	43	42	42	4.7	4.4	100		<u>8</u> 5	47		40		99		84	48	47	0.8	Her3 86	2
		44	46	63			62	46	60	57	60	62	60	60	61	47	61	63	45	76	75	62	61	59	60	62	59	46	61	49	62	86	20	49	46	50	46	76	74	48	87	
	85		84	49		85	49	85	46	46	47	49	45	49	45	86	45	51	83	52	52	49	45	44	43	49	46	86	47	95	49	50	49	84	85		85		51		88	
		43	47	64	73	47	63		58		59				0.9	47	60	70	46	75	74	66	60	58	58	65	61	47	62	48	66	75	80	49	47	49	47	75	73	46	89	$\frac{1}{2}$
		40	44	59	67		72	43	100	88	99	61	99	61	100	43	100	61		61	60	61	100	91			58		62			63			43		44		59		90	$\downarrow$
וש	85	83	84	49		85	49	85	46		47	49	45	49	45		45	51	83	52	52	49	45	44	43	49	46		47		49	50	49		85		8.5			85	91	
FUDIX		43	47	64	73	47	63	47	58	57	68	66	59	60	60	47	09	70	46	75	74				58		61	47	62	48	99	75	8		47		47	75	73	46	92	4
1	84	95	94	47	48	79	47	98	44	43	45	47	43	48	43	98	43	49	82	48		47	43	42	42	47	44	99	47	36	47	48	48	9	98	8	84	48	47	79	93	
	83	91	94	49	50		49	94	45		46	49	44	49	44	4	44	50	_	50	50	┝╌	$\vdash$		42	8	46	5	49	4	49	의	48	4	4	-	3	50		82	94	-
3		42	46	81	69	49	81				62		62		63	5	63	7	4	77	6	2	3	0	1	4	88	5	81	7	$\dashv$	7	66	0	45	1	45	77	75	48	95	$\frac{1}{1}$
2	47	43	47	64	73	47	63	47	58	57	59		59	60	60	7	0 9	0	$\vdash$	75			0	8	58	5		7	62	8	99	75	-	$\dashv$	47		Н		-	46	96	$\frac{1}{1}$
	83	95	94	47	48	79	47	86	44	43	45	47	43	48	43	98	43	49	81	48	48		43	42	42	48	44	99	47		47	48	49		_	78	83	48	47	79 1	97	$\frac{1}{2}$
	81	78	79		51	100	50		46	44	46	49	44	48	44	80	44	Н	9	0	0	9	┢	ω	2	8	-	0	5	0	-	9	6	ω		3	8 1	0	49	00 1	9 8	4
	81	78	79		51	100	0	╁─		44	46	49	44		-	$\vdash$	4	9		0	0	9	4	3	2	┢	6	$\vdash$	45	0	9	9	$\dashv$	ω	_	-	81	50	49	00	99 1	4
	_	8	9	9	51	0	-	9	46	44	46	-	4	8	$\vdash$	-	44	$\vdash$	9	0	⊢	9	4	3	-	8	-	0	5	0	-	9	$\dashv$	ω	9	-	$\vdash$	0	49	00	의	$\frac{1}{2}$
					8			97		3	5	47		8	43	97	43	4 9	81	8	8	7	3	42	42	47	44	98	46	8.4	47	47	48	78	97	77	83	48	47	79	<u>=</u>	5
		. Jn	1 4 h	: =	iæ	10	17	98	4	10	15	17	įΰ	ι &	iω	-		<u> </u>	· —	-	-	i		<del>-</del>	i	•	-	-	17			_	_				-	ا مُا			7	
	 	78		1	5	-	1	9	1	1-	46	9	1	8		0	-	9	9	0	0	9	4	ω	-	8	6	0	45	0	9	9	6	3	9	ω	_		49		03 1	4
	47	43	47		73						59					7		0	6	75   1	4	6	0	8	8	5	_	7	62	8	6	75	08	9	7		7	75   1		46	04	4
	49		49	70		L .					60		60		61	8	61	ω	48	001	9	ω	-		59			48	70	50		79	72				9	00	98	0	05 1	
	82	93	92			78	46			42	44	46	42	47			42		0.8			46	42		4		43	97		84	46	47	47	77	_	6	82	-	46	8	10,1	-
			95				47		44	43	45				43	99	43		82			47				47	44	00	47	85		48	48	79	99		84			80	107	

APPENDIX ++= 7264

Γ	Ţ	Т	. [	45		5				6	4		6		ر ن	6	46		6			9	46	45	46	59	48		46	46	46	5	7	99	98	73	46	64	4		63	46
-	+	+	4	5 8	9	9	4	4	ဖ	4	ᅴ	4	4	7 4	4	-	-	<del>-</del>		6 46	9 47	9 47	æ	5 46	4	9.44	-	5	8	<u>ω</u>	80	7 4 4	4	4	4	4	ဖ	4	8	8	4	5
-	'	<u> </u>	6	=	7 4	4	9 4	9 5		6 4	8	4	4	7 4	4	4	8	7	4	-	1	4	<u>@</u>	4	4	4		<u>의</u> 8	7	<u></u>	7		4	4	4	4	<u></u>	5	8	<u></u>	4	0 8
	-	╁	┪	00 8	46 5	4	9	1 5	<u>-</u>	9 4	-	5	7 4	4	4		<u>_</u>	7	8 4	4	5		0	4	4	4	-1 -8	8	8	0	9	7 4	4		4	4	8	5	8	8	9 4	
	n C	+	9	5	8	5 6	2 9	<u>-</u>	4	$\dashv$	5 4	7 7	7	7	5 7	7	5 4	4	8 7	7	0	$\dashv$	6 49	-	8	6	5	5 4	4	4	4	6	7	8	8	6	4	7	5	4		4
1	- 4	١		1 4	0 6	5			$\dashv$	8	$\dashv$	5 7	4 7	6 7	2 7		-	4	5 7	7	0	5	4	4	4	6	$\dashv$	9 45	$\dashv$	9 47	4		6	6	5	5	4	7	4	4		4
		7	9	7	0	$\dashv$	7	0	4	의	5		8	4 4		$\dashv$	$\dashv$	$\dashv$	9 4	-	$\dashv$	9 47	7 10	5 44		0 43	$\dashv$	5 84	$\dashv$	_	$\dashv$	8 46		$\dashv$	9 46	7 43	5 8			_		<u></u>
				0 4	$\vdash$	43 6	$\dashv$	9 7	$\vdash$	47 6	$\dashv$	6	$\neg$	8	-	47 6	7	$\dashv$	46 6	47 6	47 7	7 7	00 4	-	6 4	ය 6	7 4	4	$\dashv$	러	9	$\dashv$	5 7	7 7	$\dashv$	3 6	3 4	8	00 4	4	9	3 4
0	0 0	2	75	8	76 5	_		_	6	2	6	2	0	2	8	2	$\dashv$	41 7	_	2	6 5	-	6	5	4	_	6	8	-	46 8	6		-+	6		8	$\dashv$	<del>-</del> 5	$\dashv$	6 8	3 4	7 8
	ח כ	7	9	5	0 1	5	2	_	-	49 6	$\dashv$	7 6	-			9 6	-	$\neg$	8	8	의	0 1	Н		48 4	-	의	5	5 4	್ರ	5	7 5	6	<u> </u>	9 9	4 7		0	6 4	6 4	8 6	4
$\vdash$	ר מ	4	99	6	]	_	79   5	$\dashv$	6	6	7	36 5	_	68 6	$\vdash$	6   6	$\dashv$	$\dashv$	5		$\Box$	00  5	47 4	-	7 4	60 1	49 4	7 4	7 4	47 4	7	8	$\neg$	9 00		4 5	7 4	5 6	8 4	7 4	4 6	7 4
1	1	4	59	4	0 0	100	9	$\vdash$	43	_	4	9	9	1	4	1	$\vdash$	-	0	0	0	9	3	8	9	00 4	46 8	4 8	2	з 8	2 8	7 4	_	0	9	4	4	1 5	4	ယ 8		4 8
		+	ဖ	5	50 1	5	52	_		H	-	-	_	49		_	8.5	_		_	-		-	_	48	_	08	5	5	6	5	7	6	0	9	44 7	34 4	6   0 i	6	6	8 8	4
0	2 1		99	46	00	60	9			66	7		4			-	47	42				100	-	46	-				47	47	7			100	9	74	7	5	8	7	34 4	17 8
	70		46	79	47	43	48	49	83	7		46	Н	48			$\vdash$	74	46	47	H		$\vdash$	44	46		-	Н	8	99		-			Н	43	83	48	99	99	6	ω
		83	47	82	48	44	49	51	82		83	Н	47	49	$\vdash$	49	$\vdash$	73	48	$\vdash$	48	-	95	-	46	$\vdash$	79	$\vdash$	$\vdash$	95	$\vdash$	47	$\vdash$	48			82	50		95		82
		4.5	71	48	72	63	77	77	44		45	81	90	83	91	92	45	41	91	82	72	⊢	┢╌	┢	$\vdash$	-	48	⊢				-	65		-	63	45	90	47	45		46
` {_	٦).	7	99	46	100	60	9		6	99	7	6	4	68	0	6	7	42		67	ŏ	_	<b>!</b>	46	<b>├</b>	0	9	_		47		$\vdash$	$\vdash$	100	-	74	-		48	47	64	47
, 9	70	8 3	46	79	47	43	48	50	82	47	83	46	45	48	43	47	83	74	46	47	47	47	99	44	46	43	76	83	98	99	98	46	45	47	46	43	82	48	99	99	46	82
0		2	45	100	46	44	49	5	81	49	81	45	47	47	46	49	8 1	72	48	46	46	46	80	45	45	44	8 1	8 1	79	80	79	47	47	46	45	43	80	50	80	80		8 1
9			45	100	46	44	49	51	81	49	81	45	47	47	46	49	81	72	48	46	46	46	80	45	45	44	8 1	81	79	80	79	47	47	46	45	43	80	50	80	80		81
6			45	100	46	44	49	51	81	49	81	45	47	47	46	49	81	72	48	46	46	46	80	45	45	44	81	8 1	79	80	79	47	47	46		43		50		80		81
		83	46	79	47	43	48	40	82	47	83	46	45	48	43	47	83	73	46	47		47	98	45	47		77		97		97		45	47	46	43		48		98		82
-		œ သ	46			:	φ (Φ)		82	4	3		් <u>ය</u> ුපා	4	44	44	3	-	4.	15	1	4	99	4	5.	4	-1	သိ	9 8	9.9	98	40	<u> </u>	14	14	1.	ļ∝	\$ 63	9	φ	÷	α
		8	400	100	46	44	49	51	81	49	8-1	45	47	47	16	19	32. -	12	8	6	46	46	80	5	÷5	4	8	8 1		80	79	47	17	6	5	ώ	30	0	30	0	9	
		47	99	46	100	60	79	67	46	66	47	66	64	68	60	99	47	42	65	67	100	100	4	46	47	60		47		47	47			100	99	74	47			47	64	47
	50	49		50	75			85	48		49							43		73		75		46			51	49	48	48	48	65	70	75	74		49		49	48	70	49
	78		45	78	46	42			8-1	46		45			42	46	82	72	45	46	46	46	9 /	43			75		96				44						97	97	45	8 1
	80	84	46	80		43		49			84		45	48	43	4 /	84	7.4	46		4.7	4 /	100	44	46	43	77	84	99	100	99	46	45	47	46	43	83	48	100	100	46	

APPENDEX -12- 6.18

	80																			* .																		
	81														Ī																							
	82																																					
	83							T						Ī						Ì		Ì																
	84						-	Ť	1		_			ľ	$\dagger$	1	1												1				Ĭ,					53
	85							t	1				-	-	$\dagger$	1	1	1				1	1	_			1		1								89	48
Her3	86	<u> </u>						1	1		-		1	$\dagger$	†	1	7	1		7		1	7	1		1	1	1	7		7					47	49	86
3	87	_	_			-	l	t	1				İ	İ		1				1									1						46	59	78	47
-	88	$\vdash$			-		<u> </u> -	$\dagger$	1		-	-	-	l	+	+													1				•	47	86	48	53	100
	89	┝			-	$\vdash$	$\mid$	$\dagger$	1	_	-			-	$\dagger$	1				$\dashv$	$\exists$				T								50	76	47	<del>├</del>	80	М
-	90				-	-	$\dagger$	$\dagger$	-		-	-	1		+	$\dashv$	$\dashv$	_	$\dashv$			1	$\dashv$									60	45	61	43	6	6	4
	91	-	$\vdash$	_	-	$\vdash$	-	+			$\mid$	-	$\dagger$	+	$\dagger$	1	1	ㅓ	-										$\dashv$		45	50	100	47	86	$\vdash$	53	100
-	1 92	-	-	$\vdash$				$\dagger$	-		H		╁	$\mid$	$\frac{1}{1}$	1	-												-	50	_	100	0 50	76	47	60		
-	6		-	-	-		$\vdash$	$\dagger$	1		-	-	+	╀	$\dagger$	$\dashv$	-	_		·			1						47	-	43	0 47	87	3 46	99	$\vdash$	49	87
$\vdash$	3 94	1-			-		ŀ	$\dagger$	-		┢	╁	╁	+	$\dagger$	-	-	-										94	48	85	-	48	$\vdash$	48	95	$\vdash$	$\vdash$	85
$\vdash$	9		-	-	-	$\vdash$	+	+		-	-	$\vdash$		+	+	1	_							_		-	47	1 45	3 72	-	63	-	┥	3 65	$\vdash$	11	82	Н
-	5 9	┢		-	-	-	+	+	_		-	-	+	+	+		-			_						72	7 48	5 47	2 100	50	$\overline{}$	100	50	-	47		9 80	50
-	6 97	╂─	$\vdash$	-	-	-	+	+	1		-	+	+	+	+	-			_				-		47	2 45	$\dashv$	7 98	$\neg$			_	85	$\vdash$	-	$\vdash$	) 49	85
-		┞		-	-	$\frac{1}{1}$	-	+		_	-	+	+	+	+	-				٠.١			-	79	Н	$\vdash$	$\vdash$	-	-	5   85	-	7 46	┼	-	├—	╁╌┤	H	5 85
$\vdash$	98 9	╂╌	-	$\vdash$		$\vdash$	╁	+		_	-	-	+	-	+	-				6, 3			Щ	-	Н	8 48	$\vdash$	Н	-	5 85	_		5 85	⊢	₩	$\vdash$	-	5 85
-	99 10	₩.	-	-	  -	$\vdash$	+	+	_	L	-	-	╀	+	+	-			_	11 4.75 F		. 100	100 100	_	Н	8 48	-	$\vdash$	6 46	5   85	-	6 46	┼─	┼	<del> </del>	<del> </del>	1 51	╅╼┪
6	1001	_	$\vdash$	$\vdash$	-	+	+	+	4	_	┞		+	+	+	_		_		ς λ	. 79	0 79	0 79	9 97	-	-	-	-		-	4   43	6 47	┼-	-	<del>-</del>	1-	1 49	╁─┤
9	01	1		1_	_	1 -		_	_	_	-	ļ_	-	+	_	_		_		30	⊢	H	$\vdash$	$\vdash$		5 45	-	7 98	$\vdash$	_	3 43	$\vdash$	5 86	╅─	<del> </del> —	$\vdash$	9	5
	1-			• · · ·			•	•	1 49-	•	• Î	+	· • · · ·					_				-			•—	•	•—-	•		-	-	•	•	•—	•—	•	. — . —:	13.
-	1031	1	-	-	-	-	+	$\frac{1}{1}$	_	-	┞	+	+	+	+			┢╌	80 4	-	i	100		⊢		48 7	-		-	⊢	┢	ļ	85 5	┼	╁	+	<del>i</del> —	1-1
-	04	1	-	-	-	-	+	-			-	+	1	+			-	46 5	7	47 :4	46 5	╌	├-	-	ĭ	₩			100 7	_	60 6	1	50 5	t-	7	╁─	80 9	╁╌┪
	105	1-	-	-	-	+-	-	-		-	-	-	-	+	-		75	0	↓	8	0	0		48	╀	77 '	<b>↓</b>	-	-	2	_	5	2	6	-	┿	4	2
-	106	1-	-	1	-	_	+	-		_	-	-	-	+	·	47	_	•	┪	<del> </del>	78	<del>† -</del>	<del>† –</del>	_	1-	1-	┼-	96	┥	┝	42	┼	85	+	<b>∔</b> —	↓_	48	<del> </del>
	c >												1.		97	48	47	80	99	86	80	08	80	99	47	45	95	99	47	86	43	47	86	46	100	47	49	86

Mpl.14		46	48	44	84	84	85	46	44
3 3	3	43	44	40	96		83	43	40
4 3			46	44	95	95	84	48	44
9	50	84	65	59	47	47			59
51 36	54	66	76	66	48	48	51	66	67
5 3	5		49		80	80	85	50	45
9 3	0		67	72	47	47	49	78	72
5 3	5		46	43		99	85	47	43
6 3	46 4		60	99	44	44	46	_	100
ပ	5	57	55		43	43	46	56	88
47 30	6		60	98	45	45	47	59	99
2	Н	100	66	61	47	47	49	99	61
2	6	60	6	98	43	43	45	59	99
$\dashv$	_	80	62		48		49	80	61
2	9	61	6	99	43		45	60	9
2	5		46		99	66	98	47	43
45 24	46 4	61	61	99	43		45	09	100
2	2	94	69	61	49	49	51	94	61
3	$\vdash$	45	47	43	82	82	83	45	43
-	3	73	71	60	48	48	52	72	61
Н	3	72	70	59	48	48	52	71	60
-	0	100	66	61	47	47	49	99	61
Н	6	61	61	99	43	43	45	60	9
Н	5	58	57	90	42	42	44	57	91
43 16	4	59	59	97	42	42	43	58	98
$\dashv$	-	90	66	62	47	47	49	89	62
6 14	7	95	62	58	44	44	46	94	58
$\dashv$		47	46		90	100	86	47	43
Н	8	85	65	62	47	47	47	84	62
		48	50		85	85	95	48	45
9 10	0	100	66		47	47	49	99	61
	_	66	72		48	48	50	66	63
	-	64	77	58	48	48	49	64	58
4	3	52	50		79	79	84	52	48
5	⊢	47	46	43	99	99	85		43
ω		52	51	48	78	78	83	52	48
5	83 8	46	48	44	84	84	85	46	44
-	-	73	71		48		52	72	61
1 2	2 5	71	70	58	47	47		70	59
5 1	8	49	48	44	80	80	85	49	44
17	116 11	115	114	113	112	111	110	109	108
	_			VEGF					

APPENDIX -14- 6.14

<del>·</del> <del>·</del> <del>·</del> <del>·</del> <del>·</del> <del>·</del> <del>·</del> <del>·</del> <del>·</del> <del>·</del> <del>·</del>	85 84 49 47 49 48 47 46 85 84 47 46 85 84 47 46 85 84 47 46 85 84 47 46 85 84 47 46 85 84 47 46 85 84 85 84 85 43	75 74 85 84 49 47 49 48 47 46 85 84 47 46 85 84 85 84 85 84 87 49 87 40 87 40	88 48 47 98 48 46 41 75 74 46 85 84 99 49 47 94 45 43 90 49 48 97 47 45 88 47 46 46 85 84 99 49 47 46 85 84 90 49 47 47 46 48 85 84 90 49 47 48 83 82 51 49 72 52 48 60 45 43 60 45 43 60 45 86	66 50 47 88 48 47 98 48 46 41 75 74 46 85 84 90 49 49 97 47 45 88 47 46 85 84 99 49 48 97 47 45 86 85 84 98 85 84 98 85 84 98 85 84 98 85 84 98 85 84	47 86 100 66 50 47 66 50 47 88 48 47 98 48 47 41 75 74 46 85 84 90 49 47 91 45 43 90 49 48 97 47 46 88 47 46 88 47 46 88 47 46 88 47 46 85 84 87 47 46 87 87 88 88 47 46 88 49 47 46 80 49 49 47 40 48 83 66 50 47 46 85 84	48 48 48 46 47 46 47 86 100 66 50 47 88 48 47 46 47 46 47 46 47 46 47 45 43 45 45 84 85 84 85 84 85 86 50 47 45 85 84 85 86 50 45 45 85 86 50 45 45 45 45 85 86 50 45 45 45 45 85 86 50 45 45 45 85 86 50 45 45 45 45 45 45 45 45 45 45 45 45 45	48       48       48       46         47       46       46       44         47       86       100       47         66       50       47       46         98       48       48       47         46       85       84       46         99       49       49       47         90       49       47       45         46       85       84       47         99       49       47       45         46       85       84       83         99       49       47       46         88       47       46       85         84       87       46       83         99       49       47       45         48       47       46       85         84       82       51       49         45       43       83         0       66       50       47         46       85       80	40 80 77  49 80 77  48 48 48 46  47 46 44  47 86 50 47  88 48 48 47  46 85 84  99 49 47 46  46 85 84  97 47 46  48 85 84  99 49 47  99 49 47  99 49 47  90 60 45 43  0 60 45 43  0 60 45 43  0 60 45 43  0 60 45 43	47 85 99 46 85 84 49 80 77 960 45 43 47 46 44 47 46 44 47 86 100 66 50 47 88 48 47 46 85 84 90 49 47 94 45 43 90 49 47 46 85 84 97 47 46 46 85 84 97 47 45 98 48 47 99 49 47 99 49 48 97 47 45 88 47 46 98 85 84 99 49 47 99 49 48 97 47 45 88 47 46 98 85 84 99 49 47 99 49 48	47 86 100 47 85 99 46 85 84 49 80 77 60 45 43 48 48 46 47 46 44 47 86 100 66 50 47 88 48 47 46 85 84 47 46 85 84 90 49 49 47 91 45 43 92 49 47 93 49 47 94 45 43 90 60 45 84 97 47 46 88 85 84 97 47 46 88 85 84 97 47 46 88 85 84 97 47 46 88 85 84 99 49 47 99 49 48	2 47 85 99 2 47 86 100 2 47 86 100 2 47 86 99 4 46 85 84 6 49 80 77 6 49 86 50 47 0 66 50 47 0 98 48 48 47 46 85 0 47 46 85 0 47 46 44 1 99 49 48 1 99 49 47 1 99 49 48 1 99 49	1 65 46 45 7 78 47 46 7 78 47 46 2 47 85 99 2 47 85 99 4 46 85 84 4 46 85 84 6 49 80 77 6 49 86 100 9 66 50 47 9 66 50 47 0 98 48 48 1 99 49 47 1 4 82 51 49 1 4 9 85 84 1 4 9 85 84 1 4 9 85 84 1 4 9 85 84 1 9 9 72 52 48 1 4 9 85 80 1 4 9 85 84 1 4 83 1 4 9 85 80 1 4 9 85 84 1 4 82 51 49 1 4 9 85 84 1 4 9 85 84 1 4 9 85 84 1 4 9 4 83 1 4 9 85 84 1 4 9 85 84 1 4 9 85 84 1 4 9 85 84 1 4 9 85 84	0 66 50 47 1 65 46 45 7 78 47 46 2 47 85 99 2 47 86 100 2 47 86 99 4 46 85 84 6 49 80 77 8 47 86 100 9 66 50 45 9 66 50 47 8 48 48 47 9 66 50 47 10 66 85 84 10 49 49 47 10 49 48 48 10 49 49 47 10 49 48 10 49 49 47 10 49 49 47 10 60 45 43 10 60 45 43	9 65 49 46 1 65 46 45 1 65 46 45 1 7 78 47 46 2 47 85 99 4 46 85 84 6 49 80 77 0 60 45 43 0 66 50 47 0 88 48 47 0 46 85 84 1 99 49 47 1 99 49 49 47 1 99 49 47 1 99 49 47 1 99 49 47 1 99 49 47 1 99 49 49 47 1 99 49 49 47 1 99 49 49 47 1 99 49 49 47 1 99 49 49 47 1 99 49 49 47 1 99 49 49 47 1 99 49 49 47 1 99 49 49 47 1 99 49 49 47 1 99 49 49 47 1 99 49 49 47 1 99 49 49 47 1 99 49 49 49 49 49 49 49 49 49 49 49 49	65 49 46 66 50 47 65 46 45 78 47 46 47 85 99 46 85 84 47 86 100 47 86 100 66 50 47 66 50 47 66 50 47 98 48 48 46 45 47 86 100 68 50 47 98 48 48 47 86 100 68 50 47 99 49 49 47 90 49 49 47 90 49 49 47 91 45 84 92 49 47 93 45 84 94 45 84 95 84 96 85 84 97 47 46 98 85 84 99 49 47 99 49 47 99 49 47 99 49 47 90 60 45 84 90 60 45 84	65 46 44 43 65 46 45 47 86 50 47 46 47 86 100 47 46 47 86 100 47 46 47 86 50 47 46 46 47 86 50 47 46 46 47 86 50 47 46 85 84 47 46 47 46 85 84 47 46 85 84 47 46 85 84 47 46 85 84 47 46 85 84 47 46 85 84 47 46 85 84 83 0 66 50 45 43 0 66 50	1 96 50 48 4 46 84 83 9 65 49 46 7 78 47 86 7 78 47 86 9 66 49 80 77 0 66 49 80 77 0 66 49 80 77 0 66 49 80 77 0 66 49 80 77 0 66 50 47 0 98 48 48 46 1 90 49 48 47 1 90 49 48 48 1 90 49 48 85 1 90 49 4	96 50 48 49 46 45 49 46 45 49 46 45 49 46 45 49 46 45 49 46 45 49 46 45 49 46 46 45 49 46 46 45 46 45 46 46 45 46 46 45 46 46 46 46 46 46 46 46 46 46 46 46 46	47 86 100 48 86 100 48 86 100 48 86 100 46 84 83 61 44 43 65 49 46 47 85 99 47 86 100 47 86 100 47 86 100 66 50 47 66 85 84 48 48 46 49 49 49 49 49 49 49 49 40 88 48 48 41 75 74 42 88 48 48 43 99 49 49 47 44 85 84 46 85 84 47 86 100 66 50 47 66 50 47 66 85 84 47 46 85 84 48 48 46 49 49 49 49 47 46 85 84 47 46 85 84 48 48 47 48 85 84 49 49 49 47 40 88 85 84 41 42 86 85 84 42 48 85 84 43 99 49 49 47 46 85 84 47 46 85 84 48 85 84 48 85 84 48 85 84 48 85 84 48 85 84 48 85 84 48 85 84 48 85 84 48 85 84 48 85 84 48 85 84 48 85 84 48 85 84 49 47 46 40 48 85 84 41 42 85 84 42 48 85 84 85 84 45 84 85 86 85 84 47 46 85 84 48 85 84 48 85 84 48 85 84 48 85 84 48 85 84 48 85 84 48 85 84 48 85 84 48 85 84 49 47 46 48 85 84 49 47 46 49 47 48
80 44 43 80 80 80 80 80 80 80 80 80 80 80 80 80	85 84 49 47 49 48 47 46 85 84 47 46 85 84 47 48 50 47 85 80	75 74 85 84 49 47 49 48 47 45 85 84 47 46 85 84 87 49 87 49 87 49 85 84	48 47 48 46 75 74 85 84 49 47 49 48 47 45 85 84 47 46 85 84 87 49 87 49 89 49	50 47 48 47 48 46 75 74 85 84 47 45 47 45 47 45 47 45 87 84 48 47 49 48 40 47 40 48 40 47 40 48 40 47 40 48 40 47 40 48 40 47 40 48 40 47 40 48 40 47 40 48 40 47 40 48 40 47 40 48 40 47 40 48 40 47 40 48 40 47 40 48 40 47 40 47 40 48 40 47 40 48 40 47 40 48 40 47 40 40 47 40 40 47 40 47 40 47 40 47 40 47 40 47 40 47 40 47 40 47 40 47 40 47 40 47 40 47 40 47 40 47 40 47 40 47 40 47 40	86 100 1 50 47 48 47 48 46 75 74 85 84 49 47 49 48 47 46 85 84 47 46 85 84 47 46 85 84 47 46 85 84 47 46 85 84 47 46 85 84 47 46 85 84 85 84 87 86 87 86 88 86	48 46 44 46 46 47 48 46 47 48 46 47 48 46 47 49 47 48 83 84 83 84 83 84 83 84 83 84 83 84 83 84 83 84 83 84 83 84 83 85 86 86 86 86 86 86 86 86 86 86 86 86 86	45 43 46 46 47 48 47 48 47 45 43 44 46 47 45 43 47 46 85 84 83 50 47 45 43 45 43 45 45 45 45 45 45 45 45 45 45 45 45 45	80 77 48 46 44 4 48 47 48 48 47 48 48 47 48 47 46 85 84 47 46 85 84 83 85 86 87 85 86 87 85 86 85 86 85 86 85 86 85 86 85 86 86 85 86 86 85 86 86 86 86 86 86 86 86 86 86 86 86 86	85 99 0 85 84 1 80 77 1 45 43 4 86 100 1 50 47 1 48 46 4 48 47 4 85 84 4 85 84 4 85 84 83 5 50 47 46 83 85 84 83 85 86 85 80	86 100 1 85 99 9 86 77 8 48 46 44 4 46 44 4 86 100 1 86 75 74 4 88 84 46 47 46 47 45 43 49 47 45 43 49 47 45 43 48 83 84 83 85 86 85 85 85 85 85 85 85 85 85 85 85 85 85	85 99 9 9 9 9 9 9 9 9 9 9 9 9 9 9 9 9 9	46 45 47 46 45 85 99 6 85 99 6 85 84 46 44 4 46 44 4 50 47 4 48 46 4 48 46 4 48 46 4 48 46 4 48 46 4 48 46 4 48 47 4 49 47 4 49 47 4 49 47 4 49 47 4 49 47 4 50 47 4 60 47 48 4 60 47 4 60 47 4 60 47 4 60 47 4 60 47 4 60 47 4 60 47 4 60 47 4 60 47 4 60 47 4 60 47 4 60 47 4 60 47 4 60 47 48 4 60 47 48 4 60 47 4	5 46 47 46 48 48 47 46 85 84 47 46 47 46 85 84 47 46 85 84 83 86 650 47 46 85 84 83 85 84 83 85 84 83 85 84 83 85 84 83 85 85 84 83 85 85 85 85 85 85 85 85 85 85 85 85 85	5 49 46 45 49 46 46 6 85 84 84 46 47 46 85 84 84 84 84 84 84 84 84 84 84 84 84 84	1 1 44 43 4 5 49 46 47 46 47 7 85 99 9 9 9 9 9 9 9 9 9 9 9 9 9 9 9 9 9	6 84 83 8 6 84 43 4 7 85 99 9 7 86 100 1 7 86 100 1 7 86 99 9 8 847 46 45 43 8 8 48 48 46 44 8 7 86 100 1 8 84 84 84 8 8 84 84 8 8 85 84 8 8 85 84 8 8 85 84 8 8 85 84 8 8 85 85 8 8 85 85 85 8 8 85 85 85 85 85 85 85 85 85 85 85 85 85	6 50 48 48 46 44 49 46 49 46 49 49 49 49 49 49 49 49 49 49 49 49 49	8 8 6 100 1 6 8 4 8 3 8 6 8 4 8 3 8 6 8 4 8 3 8 7 8 5 49 46 4 7 8 5 99 9 7 8 6 100 1 7 8 7 8 7 8 8 8 8 8 8 8 8 8 8 8 8 8 8 8 8 8 8 8	7 86 100 1 8 86 100 1 8 86 100 1 8 86 100 1 8 86 100 1 8 87 49 46 43 46 45 49 46 45 49 46 45 49 46 45 49 46 47 46 47 47 47 47 47 47 47 47 47 47 47 47 47
444444444444444444444444444444444444444	85 84 49 47 49 48 47 46 85 84 47 46 85 84 47 46 87 88 50 47	75 74 85 84 49 47 49 48 47 45 85 84 47 46 85 84 85 84 87 49 47 46 49 47 49 48 49 47 49 47 49 48	48 47 48 46 75 74 85 84 49 47 49 48 47 45 85 84 47 46 85 84 85 84 85 84 87 49 50 47	50 47 48 47 48 46 75 74 85 84 47 45 47 45 84 83 84 83 85 84 87 49 88 84 88	86 100 1 50 47 48 47 48 46 75 74 85 84 49 47 49 48 47 46 85 84 47 46 85 84 47 46 85 84 47 46 85 84 47 46 85 84 47 46 85 84 47 46 85 84	48 46 44 46 46 47 48 46 47 48 46 47 48 46 47 49 47 46 85 84 47 46 85 84 83 85 85 85 85 85 85 85 85 85 85 85 85 85	45 43 46 46 47 48 46 47 48 46 47 48 47 45 43 47 45 43 45 45 43 45 45 45 45 45 45 45 45 45 45 45 45 45	80 77 48 46 44 4 86 100 1 86 100 1 86 100 1 86 100 1 85 84 47 45 84 85 85 85 85 85 85 85 85 85 85 85 85 85	85 99 0 85 84 1 80 77 1 45 43 4 46 46 4 48 100 1 50 47 4 48 47 4 49 47 45 84 4 49 47 45 84 83 84 83 85 84 83 85 84 83 85 84 83 87 87 87 87 87 87 87 87 87 87 87 87 87	86 100 1 85 99 9 86 77 1 48 46 44 1 50 47 1 48 46 100 1 50 47 1 48 46 47 1 48 46 47 1 49 48 46 1 47 45 43 1 85 84 47 1 49 48 46 1 47 45 43 1 85 84 83 1 50 47 1 49 48 46 1 47 45 1 49 48 46 1 47 45 1 49 48 1 50 47 1 49 48 1 40 49 47 1 40 49 48 1 50 47 1 40 49 48 1 50 47 1 40 48 1 40 49 47 1 40 49 48 1 50 47 1 40 48 1 40 49 48 1 50 47 1 40 48 1 40 48 1 40 48 1 40 48 1 40 48 1 40 48 1 40 48 1 40 48 1 40 48 1 40 48 1 40 48 1 40 48 1 40 48 1 40 48 1 40 48 1 40 47 1 40 48 1 40 40 1 40 1 4	85 99 9 9 9 9 9 9 9 9 9 9 9 9 9 9 9 9 9	46 45 47 46 85 99 6 85 99 6 85 84 48 46 48 46 48 46 48 47 48 46 49 47 49 48 49 47 49 47 49 48 49 47 49 48 49 47 49 48 49 47 49 48 49 49 49 49 49 49 49 49 49 49 49 49 49 49 40 40 40 40	50 47 46 47 46 48 47 45 48 47 46 48 47 46 48 47 46 48 47 46 48 47 46 48 47 46 48 47 46 48 47 46 48 47 46 48 47 46 48 47 46 48 47 46 48 47 46 48 47 46 48 47 46 48 47 46 47 47 46 47 46 47 46 47 46 47 46 47 46 47 46 47 46 47 46 47 46 47 47 46 47 47 46 47 47 46 47 47 46 47 47 46 47 47 46 47 47 46 47 47 46 47 47 46 47 47 46 47 47 46 47 47 46 47 47 47 46 47 47 46 47 47 46 47 47 46 47 47 46 47 47 46 47 47 46 47 47 47 47 47 47 47 47 47 47 47 47 47	5 49 46 45 49 46 46 6 85 84 8 47 46 6 85 84 8 47 86 100 1 9 9 9 9 9 9 9 9 9 9 9 9 9 9 9 9 9	1     1     44     43       5     49     46     46       6     50     47     46       7     85     99     6       7     85     99     6       8     48     46     45       9     9     6     85     84       100     1     46     46     46       100     1     46     46     46       100     1     46     46     46       100     1     47     46     47       100     1     47     46     47       100     1     49     47     46       100     1     49     47     46       100     1     49     48     47       100     1     49     48     47       100     1     49     48     47       100     1     49     48     46       100     1     49     48     46       100     1     49     48     46       100     1     49     48     46       100     1     49     49     49       100     1     49 <td< td=""><td>6 84 83 8 6 84 83 8 6 84 43 4 7 85 99 9 7 86 100 1 7 84 84 84 85 8 84 84 85 8 84 85 85 8 84 85 85 8 85 85 8 85 85 8</td><td>6 6 50 48 49 46 45 48 48 48 48 48 48 48 48 48 48 48 48 48</td><td>8 8 8 6 100 1 6 50 48 4 83 8 6 84 83 8 7 85 49 46 45 4 7 85 99 9 9 80 77 46 44 43 4 9 80 77 7 46 44 43 4 7 86 50 47 4 6 8 48 48 46 4 4 4 7 8 5 8 4 4 6 4 4 4 7 8 6 8 5 8 4 7 4 6 4 4 4 6 4 4 6 4 6 4 6 4 6 4 6 4</td><td>7 86 100 1 1 8 8 8 6 100 1 1 8 8 8 6 100 1 1 8 8 8 6 100 1 1 1 1 1 1 1 1 1 1 1 1 1 1 1 1 1</td></td<>	6 84 83 8 6 84 83 8 6 84 43 4 7 85 99 9 7 86 100 1 7 84 84 84 85 8 84 84 85 8 84 85 85 8 84 85 85 8 85 85 8 85 85 8	6 6 50 48 49 46 45 48 48 48 48 48 48 48 48 48 48 48 48 48	8 8 8 6 100 1 6 50 48 4 83 8 6 84 83 8 7 85 49 46 45 4 7 85 99 9 9 80 77 46 44 43 4 9 80 77 7 46 44 43 4 7 86 50 47 4 6 8 48 48 46 4 4 4 7 8 5 8 4 4 6 4 4 4 7 8 6 8 5 8 4 7 4 6 4 4 4 6 4 4 6 4 6 4 6 4 6 4 6 4	7 86 100 1 1 8 8 8 6 100 1 1 8 8 8 6 100 1 1 8 8 8 6 100 1 1 1 1 1 1 1 1 1 1 1 1 1 1 1 1 1
╼╁╶╂┈╁╌╂╼╂╌╂╼╂	85 49 45 45 45 45 45 45 45 45 45 45 45 45 45	45 40 40 40 40 40 40 40 40 40 40	48 48 48 49 49 49 49 49 49 49	4 4 4 4 4 4 4 4 4 4 4 4 4 4 4 4 4 4 4	500 6 485 7 485 7 495 7 450 6 1	45000000000000000000000000000000000000	4 4 4 4 4 4 4 4 4 4 4 4 4 4 4 4 4 4 4	45 45 46 47 48 47 48 48 48 48 48 48 48 48 48 48	85 85 85 85 85 85 85 85 85 85	455 465 477 485 485 485 485 485 485 485 485	45 50 1 50 1 50 1 50 1 50 1 50 1 50 1 50	455 475 475 475 475 475 475 475	50 46 46 46 46 46 46 46 46 46 46	50 40 40 40 40 40 40 40 40 40 4	44 44 45 46 47 48 48 48 48 48 48 48 48 48 48	4 4 4 4 5 5 6 6 6 7 5 6 7 6 6 7 6 7 6 7 6 7 6 7	450 450 450 450 450 450 450 450	50 44 49 46 50 47 46 45 47 46 85 99 85 99 86 100 87 77 48 46 48 46 48 46 48 46 48 46 48 47 48 46 48 47 48 48 47 48 47 48 48 47 48 48 47 48 48 47 48 48 48 47 48 48 48 47 48 48 48 47 48 48 49 49 48 49 49 49 48 49 49 49 48 49 48 49 48 49 48 49 48 49 48 49 48 49 48 49 49 48 49 48 49 48 49 48 49 48 49 48 49 48 49 48 49 48 49 48 49 48 49 48 49 48 49 48 49 48 49 48 49 48 49 48 49 49 49 48 49 48 49 48 49 48 49 48 49 48 49 48 49 48 49 49 48 49 48 49 48 49 48 49 48 49 48 49 48 49 48 49 48 49 48 49 48 49 48 49 48 49 48 49 48 49 48 49 48 49 48 49 49 49 48 49 48 49 48 49 48 49 48 49 48 49 48 49 48 49 49 49 48 49 48 49 48 49 48 49 48 49 48 49 48 49 48 49 48	86 1000 86 1000 86 1000 86 1000 87 1000 88 1 83 84 1 83 84 1 43 85 100 85 100 86 100 87 100
4 4 4 4 4 4 4 4 4 4 4 4 4 4 4 4 4 4 4	85 84 49 47 49 48 47 45 47 46 85 84 49 47 51 49 52 48	75 74 85 84 49 47 49 48 47 45 47 46 85 84 47 46 85 84 84 83 84 83	48 47 48 46 75 74 85 84 49 47 49 48 47 45 85 84 47 46 85 84 87 46 88 83 84 83	50 47 48 47 48 46 75 74 85 84 49 47 49 48 47 45 85 84 47 45 85 84 85 84 85 84 85 84 85 84 85 84 85 84 85 84	86 100 50 47 50 47 48 46 75 74 85 84 49 47 49 48 47 45 47 45 47 45 85 84 47 45 87 84 88 84 89 87 89 87 89 88	48 46 46 44 50 47 50 47 48 47 48 47 48 47 49 48 47 46 47 48 49 47 49 48 47 46 47 46 48 47 49 48 47 48 48 47 49 48 49 47 49 48 40 48 41 49 48 42 48 43 49 48 44 49 48 45 49 48 46 49 48 47 46 48 49 48 48 48 48 48 49 48 48 49 48 48 49 48 48 49 48 48 49 48 48 49 48 48 49 48 48 49 48 48 49 48 48 49 48 48 49 48 48 49 48 48 49 49 48 49 48 48 49 49 48 49 49 48 4	45 43 46 44 46 100 50 47 50 47 48 46 75 74 85 84 49 47 49 48 47 45 85 84 67 47 68 47 68 48 68  80 77 45 43 46 44 86 100 50 47 48 47 48 46 75 74 48 47 49 47 49 48 47 45 85 84 47 45 87 84 88 83 84 83	85 99 85 84 80 77 45 43 46 44 86 100 50 47 48 46 75 74 48 47 49 48 47 46 87 74 88 84 88  86 100 85 99 86 77 80 77 48 46 46 44 86 100 50 47 48 46 75 74 85 84 47 45 49 47 49 47 49 48 85 84 85 84 86 84 87 74 88 84 87 84 88 84	85 99 85 99 85 99 85 84 48 46 48 46 48 46 48 46 48 46 48 46 47 48 48 46 47 46 48 46 49 47 49 48 49 48 49 47 49 48 49 49 49	46 45 47 46 85 99 86 100 85 84 48 46 48 46 46 44 46 44 46 44 47 46 48 47 48 47 49 48 47 46 85 84 47 46 85 84 47 46 85 84 47 46 85 84 47 46 86 100 47 47 48 46 47 46 87 74 48 46 47 46 88 47 49 47 49 48 49 48 40 48 41 49 48 42 48 43 46 44 46 45 43 46 47 47 46 48 46 47 46 48 46 47 46 48 47 49 48 49 48 40 48 41 49 48 42 48 43 46 44 46 45 46 47 46 48 47 49 48 49 48 40 48 41 49 48 42 48 43 46 44 46 45 46 47 46 48 47 48 48 48 48	50 47 46 45 87 46 85 99 85 99 85 84 48 46 48 46 48 46 48 46 48 47 48 46 47 48 49 47 49 48 47 46 85 84 47 46 85 84 85 84 85 84 85 84 85 84 85 84 85 84 86 100 87 74 88 84 85 84 86 100 86	49 46 47 46 47 46 85 99 85 99 85 84 48 46 48 46 48 46 48 47 48 46 47 48 48 47 48 47 49 48 47 46 49 47 49 48 49 48 49 47 49 48 49 47 49 48 49 47 49 48 49 47 49 48 49 48 49 47 49 48 49 48 49 47 49 48 49 48 49 47 49 48 49 47 49 48 49 49 49	44 43 49 46 40 47 46 45 47 46 85 99 86 100 86 77 48 46 46 44 86 100 50 47 48 46 47 46 48 47 48 46 47 48 48 46 47 48 88 47 48 46 47 48 48 46 48 47 48 48 49 48 40 47 41 48 42 48 43 46 44 46 45 43 46 47 47 48 48 46 47 46 48 47 48 48 49 49 49 48 49 48 49 48 49 48 49 48 49 48 49 48 49 48 49 49 48 49 48 49 48 49 48 49 48 49 48 49 48 49 48 49 48 49 48 49 48 49 48 49 48 49 48 49 48 49 48 49 48 49 48 49 4	84 83 49 46 49 46 47 46 85 99 86 100 88 59 47 46 48 48 48 46 48 46 48 46 48 46 48 47 48 46 48 47 48 46 47 46 86 100 50 47 48 47 48 46 47 48 88 100 50 47 48 46 47 48 48 46 47 46 48 47 48 48 49 48 40 47 41 49 42 48 43 46 44 47 45 48 46 47 47 46 48 47 48 48 47 46 48 48 47 46 48 48 47 46 48 48 48 48 47 46 48 48 48 48 48 48 48 48 48 48 49 49 49 48 49 48 49 48 49 48 49 48 49 48 49 48 49 48 49 49 48 49 48 49 48 49 48 49 48 49 48 49 48 49 48 49 48 49	50 48 44 43 46 45 47 46 85 99 86 100 88 599 86 100 87 74 86 100 86 100 86 100 86 100 86 100 87 74 88 100 88 10	86 100 1 84 83 44 43 45 47 46 100 1 85 99 85 99 85 99 85 99 85 99 85 99 85 99 85 99 85 94 85 99 85 94	86 1000 1 86 1000 1 86 1000 1 86 1000 1 87 100 1 88 1 83 1 44 4 43 1 45 46 1 46 45 1 47 46 1 48 1 48 1 48 1 49 1 48 1 49 1 48 1 49 1 48 1 49 1 48 1 49 1 48 1 49 1 48 1 49 1 48 1 49 1 48 1 49 1 48 1 49 1 48 1 49 1 48 1 49 1 48 1 49 1 48 1 49 1 48 1 49 1 48 1 49 1 48 1 49 1 49 1 48 1 49 1 49 1 48 1 49 1 49 1 48 1 49 1 49 1 48 1 49 1 49 1 48 1 49 1		
4 4 4 4 4 4 4 4 4 4 4 4 4 4 4 4 4 4 4	84 45 46 48 47 49	5 5 7 4 5 5 6 4 3 7 4 5 5 6 4 3 7 4 5 6 5 6 4 3 7 4 5 6 6 6 6 6 6 6 6 6 6 6 6 6 6 6 6 6 6	8 46 746 8 47 9 47 1 48 1 49	1 4 6 6 7 4 7 4 6 8 3 7 4 6 8 3 7 4 6 8 3 7 4 6 8 3 7 4 6 8 3 7 4 6 8 3 7 4 6 8 3 7 4 6 8 3 7 4 6 6 8 3 7 4 6 6 8 3 7 7 8 6 8 6 8 7 7 8 7 8 7 8 7 8 7 8 7 8	5 100 1 47 0 1 5 84 47 1 6 84 7 7 4 8 8 7 4 8 8 8 7 4 8 8 8 8 8 8 8 8 8 8 8 8 8 8 8 8 8 8	1 4 6 5 8 4 6 6 7 7 4 6 5 8 8 4 4 7 7 4 6 5 8 8 4 4 7 7 4 6 5 8 8 4 4 7 7 4 6 5 8 8 4 4 7 7 7 8 6 6 8 8 4 7 7 7 8 6 6 8 8 4 7 7 7 8 6 6 8 8 4 7 7 7 8 6 6 8 8 4 7 7 7 8 6 6 8 8 4 7 7 7 8 6 6 8 8 4 7 7 7 8 6 6 8 8 4 7 7 7 8 6 6 8 8 4 7 7 7 8 6 6 8 8 4 7 7 7 8 6 6 8 8 4 7 7 7 8 6 6 8 8 4 7 7 8 7 8 8 8 8 7 8 8 8 8 8 8 8 8	43 446 47 47 47 47 47 47 47 47 48 48 48 48 48 48 48 48	77 100 100 100 100 100 100 100 100 100 1	99 0 84 1 77 1 46 1 47 1 47 1 47 1 48 4 48 4 48 4 48 4 49 1 84 8 49 1 84 8 49 1 84 8 49 1 84 8 49 1 84 8 49 1 86 1 86 1 86 1 86 1 86 1 86 1 86 1 86	100 1 99 9 100 1 100 99 99 99 99 99 99 99 99 99 99 99 99 99	45 46 46 47 47 47 47 47 47 47 47 47 47	47 45 46 46 47 47 47 47 47 47 47 47 47 47 47 47 47	46 46 47 47 47 47 47 47 47 47 47 47 47 47 47	43 46 46 46 46 46 47 47 47 47 47 47 47 47 47 47 47 47 47	83 83 84 83 84 83 83 84 84 84 84 84 84 84 84 84 84	48 48 48 48 48 48 48 48 48 48	100 1 10	100 1 100 1	
<del></del>	<del>╶╽╶╽</del> ═╂═╂═╂	<del>╶╏╶╏┈╏┈╏┈╏┈╏┈╏╸╏╸╏╸╏╸╏</del>	╼╁┈╃═╃═╂┈╂═╂═╂═╂═╂═╂═╂═┼	<del>╶┧╶┩</del> ╌╃╌╂┈╂┈╂╾╂╾╂ <del>┈╏</del> ╾╂╼╇	<del>╶╎┥┥┥┥</del> ╫╫														
000040							7465837446677700463	7465837446777046374	746583744677770463744	7465837446777704637494	7465837446777706374444444	746583744677770444444444	7465837446677770665744444444444444444444444444	7465837446777704637444445566576	7465837446777704637444455655763	74658374467777046374906576533	74658374467777046374996576338	18 6 6 6 6 5 4 4 4 5 6 6 6 6 5 4 4 5 6 6 6 6	000 44 000 44
4-004	4-4-004	04-4-004	0004-4-004	200004-4-0004	ω <b>0 0 0 0 4 - 4 - 0 0 0 4</b>	000004-4-004	000000000000000000000000000000000000000	<del></del>					4 6 6 6 7 7 7 7 7 7 7 7 7 7 7 7 7 7 7 7	4 6 6 6 4 7 7 7 7 7 7 7 7 7 7 7 7 7 7 7	4 0 0 0 0 0 0 0 0 0 0 0 0 0 0 0 0 0 0 0	440000000000000000000000000000000000000	440000000000000000000000000000000000000	4 4 9 9 6 4 2 3 2 7 1 0 9 4 4 1 4	4 6 6 6 4 7 4 6 6 6 6 6 6 6 6 6 6 6 6 6
<del>             </del>								0 0 0 0 0 0 0 0 0 0 0 0 0 0 0 0 0 0 0 0											
+-+-+	╌╂╼╂╼╂┈╂╼╂╼┨																		
000	000		44880040404			44800448800404	444800448	0 1 4 4 4 8 7 7 7 4 4 8 8 7 4 7 8	887448784488848	888744888	0 0 0 0 0 0 0 0 0 0 0 0 0 0 0 0 0 0 0 0	448887448	24498887444878444	0 0 4 4 8 8 8 8 V 4 4 4 8 0 0 0 4 4 8 0 0 0 0 0 0 0 0 0 0	4004488881448	8477844888817478	2847748888744887	80084004488880V444800V448	000000440044000000000000000000000000000
0 6	0600	06030	0603099	06030991	06000005		0603099-15896		4 6 6 6 6 6 6 6 6 6 6 6 6 6 6 6 6 6 6 6	04000000000000				0-8040408080-1-0000000					
1	12 2	ורושואוי	4 4 7 8 4 2	いいさんしん			14141410110110114141C1M1412							<del>┤╶┤┈┤┈┦┈╏┈┧┈╏╶┤═┪╌╏╶┪═╏</del> ╌╏═╅═╂═╂═┼╌┞═┼═			00000058008045540810721	<del>┩┋┩┋┩┋┩╒╃╒╃╒╃╒╃╒╃╒╇╒╇╒╇╒╇╒╇╒╇╒╇╒</del>	
_		055	0 0 0 0 0		5 5 8 8 0 0 6	0 5 5 8 8 0 0 6 6 8	0 0 6 8 5	85 85 85 85	85 85 85 85 85 85 85 85 85 85 85 85 85 8	885 86 86 87 88 86 86 86 86 86 86 86 86 86 86 86 86	8 8 8 8 8 8 8 8 8 8 8 8 8 8 8 8 8 8 8	885 447 448 856 867 868 868 878 878 878 878 87	4 4 5 5 6 6 6 6 7 7 5 6 6 6 6 6 6 7 7 5 6 6 6 6	404488888444800444	4474488884448877444	8447488888444887	030991-588968345498107221	80884404888888444880844LB	0000004404400000044400000
1						5 6 6 6 6 6 6 6 6 6 6 6 6 6 6 6 6 6 6 6	5 66 6 6 6 6 6 6 6 6 6 6 6 6 6 6 6 6 6	9 5 5 8 8 0 0 6 6 5 5 7 6 6 6 6 6 6 6 6 6 6 6 6 6 6 6	5 5 6 6 6 6 6 6 6 6 6 6 6 6 6 6 6 6 6 6	5 5 6 6 6 6 6 6 6 6 6 6 6 6 6 6 6 6 6 6	5 5 6 6 6 6 6 6 6 6 6 6 6 6 6 6 6 6 6 6	5 5 6 6 6 6 6 6 6 6 6 6 6 6 6 6 6 6 6 6	0 0 0 0 0 0 0 0 0 0 0 0 0 0 0 0 0 0 0	49 48 50 49 85 52 85 55 66 62 67 65 66 66 67 66 67 66 67 67 67 67 67 67 67	44       49       49       49       46       50       47       51       85       52       85       55       48       59       48       60       62       65       66       66	84     46       44     47       49     48       50     49       46     50       47     51       85     52       86     53       48     58       48     58       48     58       48     60       60     61       75     65       66     66       67     65	1 50 45 2 84 46 2 84 46 7 44 47 7 44 47 7 44 47 7 44 47 8 46 50 8 46 50 9 47 51 8 86 53 8 86 53 8 86 53 9 48 58 6 45 57 9 48 68 9 68 68 9 68 68 9 68 68 9 68 68 9 68 68	86 44 50 45 84 46 49 48 50 49 46 50 47 51 85 52 86 53 86 53 48 58 50 61 50 62 60 62 60 62	86 43 86 44 86 44 86 44 49 48 50 49 86 50 87 50 88 55 88 55 88 55 88 55 88 65 88
	84 84 44 48 46 83 8	5 74 74 40 44 41 80 7 5 84 84 44 48 46 83 8 6 47 47 61 66 100 50 4	8     47     47     60     65     89     49     4       8     46     46     60     65     99     49     4       5     74     74     40     44     41     80     7       5     84     84     44     48     46     83     8       5     84     84     44     48     46     83     8	0     47     47     59     78     66     51     5       0     47     47     60     79     66     51     5       8     47     47     60     65     89     49     4       8     46     46     60     65     99     49     4       5     74     74     40     44     41     80     7       5     84     84     44     48     46     83     8       6     47     47     61     66     100     50     4	5     100     100     43     46     47     85       0     47     47     59     78     66     51       0     47     47     60     79     66     51       0     47     47     60     65     89     49       8     46     46     60     65     99     49       8     46     46     60     65     99     49       5     74     74     40     44     41     80       5     84     84     44     48     46     83       6     47     47     61     66     100     50	3     46     46     49     45     48     49       3     44     44     48     46     47     48       5     100     100     43     46     47     85       5     100     100     43     46     47     85       0     47     47     59     78     66     51       0     47     47     60     79     66     51       8     47     47     60     65     89     49       8     46     46     60     65     99     49       5     74     74     40     44     41     80       5     84     84     44     48     46     83       6     87     87     61     66     100     50	43     43     99     61     61     46     49       46     46     49     45     48     49     48       44     44     48     46     47     48     48       100     100     43     46     47     85     8       47     47     59     78     66     51     5       47     47     60     79     66     51     5       47     47     60     65     89     49     49       46     46     60     65     99     49     4       74     74     40     44     41     80     7       84     84     84     46     83     8       84     84     84     46     83     8	77 77 46 50 49 78 43 43 99 61 61 46 44 44 48 46 47 48 100 100 43 46 47 85 177 47 60 79 66 51 47 47 60 65 89 49 46 46 46 60 65 99 49 74 74 40 44 41 80 84 84 44 48 46 83	99 99 42 46 47 84 84 84 44 48 46 83 77 77 46 50 49 78 43 43 99 61 61 46 44 44 48 46 49 44 44 48 46 47 48 100 100 43 46 47 85 47 47 59 78 66 51 47 47 60 65 89 49 46 46 46 60 65 99 49 74 74 74 40 44 41 80 84 84 44 48 46 83	100 100 43 46 47 85 99 99 42 46 47 84 84 84 44 48 46 83 77 77 46 50 49 78 46 46 49 45 48 49 44 44 48 46 47 48 100 100 43 46 47 48 100 100 43 46 47 85 47 47 59 78 66 51 47 47 60 65 89 49 46 46 60 65 99 49 74 74 40 44 48 83 84 84 44 48 83	100 100 43 46 47 84 99 99 42 46 47 84 84 84 44 48 46 83 77 77 46 50 49 78 46 46 49 45 48 49 46 46 49 45 48 49 100 100 43 46 47 88 100 100 43 46 47 85 100 100 43 46 47 85 100 100 65 79 66 51 47 47 60 65 89 49 46 46 60 65 99 49 74 74 40 44 48 83 77 74 40 44 48 83 77 76 66 66 51	45 45 61 99 65 4 46 46 57 60 79 4 99 99 42 46 47 8 99 99 42 46 47 8 84 84 44 48 46 8 77 77 46 50 49 7 43 43 99 61 61 4 44 44 48 46 47 8 100 100 43 46 47 4 100 100 43 46 47 8 47 47 59 78 66 5 47 47 60 65 89 4 46 46 60 65 99 4 74 74 40 44 48 46 8 84 84 44 48 46 8	47 47 60 79 66 5 45 45 61 99 65 4 46 46 57 60 79 4 99 99 42 46 47 8 99 99 42 46 47 8 84 84 44 48 46 8 77 77 46 50 49 7 43 43 99 61 61 4 44 44 48 46 47 8 100 100 43 46 47 8 100 100 43 46 47 8 100 100 43 46 47 8 47 47 60 65 89 4 46 46 60 65 99 4 74 74 40 44 41 8 84 84 44 48 46 8	46 46 59 78 65 5 47 47 60 79 66 5 48 46 57 60 79 4 99 99 42 46 47 8 99 99 42 46 47 8 84 84 44 48 46 8 77 77 46 50 49 7 46 46 49 45 48 4 100 100 43 46 47 8 100 100 43 46 47 8 84 84 84 84 64 7 47 47 59 78 66 5 47 47 60 65 89 4 146 46 60 65 99 4 147 47 40 44 48 46 8 148 46 46 60 65 99 4 148 46 46 86 65 65 89 4	43 43 54 94 61 4 46 46 59 78 65 5 47 47 60 79 66 5 48 46 46 57 60 79 4 99 99 42 46 47 8 99 99 42 46 47 8 84 84 44 48 46 8 100 100 43 46 47 8 46 46 49 45 48 46 47 47 59 78 66 5 47 47 60 65 89 4 46 46 46 60 65 99 4 84 84 44 48 46 8 84 84 84 44 48 46 8 84 84 84 44 48 46 8	83     83     44     48     46     14       43     43     54     94     61     4       46     46     59     78     65     5       47     47     60     79     66     5       46     46     57     60     79     4       49     99     42     46     47     8       100     100     43     46     47     8       84     84     84     44     48     46     8       77     77     46     49     7     48     46     8       44     44     48     46     47     4       40     100     43     46     47     4       40     100     43     46     47     4       47     47     59     78     66     5       47     47     60     65     89     4       47     47     60     65     89     4       47     74     40     44     48     46     8       84     84     84     44     48     46     8       84     84     84     44     48	48     48     61     69     97     5       83     83     44     48     46     8       43     43     54     94     61     4       46     46     59     78     65     5       47     47     60     79     66     5       45     45     61     99     65     4       46     46     47     8     46     47     8       99     99     42     46     47     8       99     99     42     46     47     8       99     99     42     46     47     8       46     46     47     48     46     47     8       47     77     77     46     49     7       43     43     99     61     61     4       47     47     49     49     79     66     5       47     47     60     65     89     4       47     47     40     44     48     46     8       47     47     40     66     5     9     4       48     84     44     48     46     6<	100 100 44 48 48 8 48 48 61 69 97 5 48 48 61 69 97 5 46 46 59 78 65 5 47 47 60 79 66 5 46 46 57 60 79 4 99 99 42 46 47 8 100 100 43 46 47 8 84 84 44 48 46 8 77 77 46 50 49 7 43 43 99 61 61 4 44 44 48 46 47 8 100 100 43 46 47 8 100 100 43 46 47 8 147 47 60 65 89 4 47 47 60 65 89 4 47 47 60 65 99 4 84 84 44 48 46 8 84 84 84 44 48 46 8	100 100 43 46 47 8 100 100 43 46 47 8 48 48 48 46 49 47 47 47 60 79 66 5 47 47 47 47 47 60 65 89 47 47 47 47 60 65 89 47 47 47 47 60 65 89 47 47 47 47 60 65 89 47 48 46 46 66 65 89 47 47 47 47 60 65 89 47 47 47 47 60 65 89 47 48 46 46 47 47 47 47 60 65 89 47 47 47 47 60 65 89 47 47 47 47 40 44 48 46 86 57 60 65 89 47 47 47 47 60 65 89 47 48 46 46 60 65 99 47 47 47 47 60 65 89 47 47 47 47 60 65 89 47 47 47 47 60 65 89 47 48 46 46 60 65 99 47 47 47 47 60 65 89 47 47 47 47 60 65 89 47 48 46 46 60 65 99 47 47 47 47 60 65 89 47 47 47 60 60 65 89 47 47 47 60 60 65 89 47 47 47 60 60 65 89 47 47 47 60 65 89 47 47 47 60 60 65 89 47 47 47 60 60 65 89 47 47 47 60 60 65 89 47 47 47 60 60 65 89 47 47 47 60 60 65 89 47 47 47 60 60 65 89 47 47 47 60 60 65 89 47 47 47 60 60 65 89 47 47 47 60 60 65 89 47 47 47 60 60 65 89 47 47 47 60 60 65 89 47 47 47 60 60 65 89

108 109 110 111 112 113 114 115 5 100 100 ω 100 100 ω 100 43 5 ი ი 6<u>1</u> 99 100 1 4 8 8 6 110 vegf.1 100 obr.19 87 her3.19 115 vegi.5 114 vegf.4 113 vegf.3 112 vegf.2 109 obr.4 108 obr.3 107 obr.26 106 obr.24 105 obr.23 104 obr.22 103 obr.21 102 obr.20 101 obr.2 99 obr.18 98 obr.17 97 obr.16 96 obr.15 95 obr.14 94 obr.12 93 obr.11 88 her3.22 86 her3.18 116 vegf.6 111 vegf.10 17 vegt.8 her3.7 her3.3 her3.11 obr.1 her3.16 her3.12 her3.4 Clone

APPENDEX -15- 6.15

#### In the Claims:

Claims 1-11, 15, 19-30 and 32 have been cancelled. Claims 34-38 have been added.

Claims 12-14, 16, 31, and 33 have been amended as follows:

12. (Amended) A multispecific antibody prepared by the method [of claim 1] comprising:

(a) expressing in a host cell a first polypeptide comprising a first heavy chain variable domain, a first or second light chain variable domain, and a first multimerization domain, wherein the first and second light chain variable domains have at least 80% amino acid sequence identity, and wherein a first binding domain is formed by the first heavy chain variable domain and the first or second light chain variable domain;

(b) expressing in the host cell a second polypeptide comprising a second heavy chain variable domain, the first or the second light chain variable domain, and a second multimerization domain, wherein a second binding domain is formed by the second heavy chain variable domain and the first or second light chain variable domain, and wherein the first and second binding domains bind different antigens;

(c) allowing the first and second polypeptides to dimerize by interaction of the first and second multimerization domains to form a multispecific antibody; and

d) recovering the multispecific antibody from the host cell.

13. (Amended) A multispecific antibody comprising a first polypeptide and at least one additional polypeptide [which meet at an interface, wherein], the multispecific antibody comprising:

[(a) the first polypertide comprises a multimerization momain

that a Souther a contract and a sign of product and the contract of a first of a first of the country and

additional polypeptides comprise a common sequence] (a) the first polypeptide which comprises a first heavy chain variable domain, a first or second light chain variable domain, and a first multimerization domain, wherein the first and second light chain variable domains have at least 80% amino acid sequence identity, and wherein a first binding domain is formed by the first heavy chain variable domain and the first or second light chain variable domain; (b) the second polypeptide which comprises a second heavy chain variable domain, the first or the second light chain variable domain, and a second multimerization domain, wherein a second binding domain is formed by the second heavy chain variable domain and the first or second light chain variable domain, and wherein the first and second binding domains bind different antigens; (c) the first and second polypeptides dimerize by interaction of the first and second multimerization domains to form a multispecific antibody. 14. (Amended) The multispecific antibody of claim 13, wherein the nucleic acid encoding the first polypeptide or the nucleic acid encoding the additional polypeptide, or both, has been altered from the original nucleic acid to encode the [interface] multimerization domain or a portion thereof. 16. (Amended) The multispecific antibody of claim 14 wherein the [interface of the] multimerization domains of the first and an additional polypeptide comprise a protuberance and cavity, respectively. 31. (Amended) The multispecific antibody of claim 13 [selected from the group consisting of] wherein the antibody is anti-Ob-R/anti-HERE I THE STORES MENT A CHEEN TO SEE seek anti-HEBer, and anti-Mail ant. HEBer.

- 34. (New) The multispecific antibody of claim 13, wherein the first and second light chain variable domains have at least 90% amino acid sequence identity.
- 35. (New) The multispecific antibody of claim 13, wherein the first and second light chain variable domains have at least 95% amino acid sequence identity.
- 36. (New) The multispecific antibody of claim 13, wherein the first and second light chain variable domains have at least 98% amino acid sequence identity.
- 37. (New) The multispecific antibody of claim 13, wherein the first and second light chain variable domains have at least 99% amino acid sequence identity.
- 38. (New) The multispecific antibody of claim 13, wherein the first and second light chain variable domains have identical amino acid sequences.